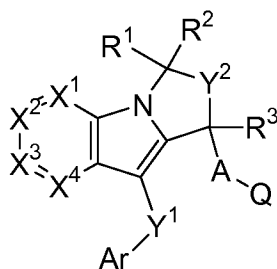


Amendment to the Claims:

Please amend Claims 1, 6, 15 and 20, and cancel Claims 3, 7 and 22-26 as follows.

Listing of Claims:

1. (Twice Amended) A compound having the formula I



I

and pharmaceutically acceptable salts and hydrates thereof, wherein:

A is selected from C₁₋₃alkyl optionally substituted with one to four halogen atoms, O(CH₂)₁₋₂, and S(CH₂)₁₋₂;

Ar is ~~aryl or heteroaryl each optionally substituted with one to four groups independently selected from R⁸~~ selected from phenyl, 2-, 3-, 4-chlorophenyl, 2-, 3-, 4-bromophenyl, 2-, 3-, 4-fluorophenyl, 3,4-dichlorophenyl, 2,3-dichlorophenyl, 2,4-dichlorophenyl, 2,5-dichlorophenyl, 2,6-dichlorophenyl, 3,5-dichlorophenyl, 3-chloro-4-fluorophenyl, 2-chloro-4-fluorophenyl, 4-chloro-2-fluorophenyl, 2-cyanophenyl, 4-methylphenyl, 4-isopropylphenyl, 4-trifluoromethylphenyl, biphenyl, naphthyl, 3-methoxyphenyl, 3-carboxyphenyl, 2-carboxamidophenyl, 4-methoxyphenyl, 3-phenoxyphenyl, 4-(4-pyridyl)phenyl, 4-methylsulfonylphenyl, 3-dimethylaminophenyl, 5-tetrazolyl, 1-methyl-5-tetrazolyl, 2-methyl-5-tetrazolyl, 2-benzothienyl, 2-benzofuranyl, 2-indolyl, 2-quinolinyl, 7-quinolinyl, 2-benzothiazolyl, 2-benzimidazolyl, 1-benzotriazolyl, 2-furanyl, 3-furanyl, 2-imidazolyl, 5-imidazolyl, 5-isoxazolyl, 4-isoxazolyl, 4-isothiazolyl, 1,2,4-oxadiazol-5-yl, 2-oxazolyl, 4-oxazolyl, 4-pyrazolyl, 5-pyrazolyl, 2-pyridyl, 3-pyridyl, 2-pyrazinyl, 5-pyrimidinyl, 2-pyrrolyl, 4-thiazolyl, 1,2,4-thiadiazol-3-yl, 1,2,5-thiadiazol-4-yl, 1,2,3-thiadiazol-4-yl, 1,2,5-oxadiazol-4-yl, 1,2,3-oxadiazol-4-yl, 1,2,4-triazol-5-yl, 1,2,3-triazol-4-yl, 3-thienyl, 1,2,4-triazol-5-yl, pyrrolopyridine, furo[3,2-b]pyridin-2-yl, thieno[2,3-b]pyridin-2-yl, 5(H)-2-oxo-4-furanyl, 5(H)-2-oxo-5-furanyl, (1H,4H)-5-oxo-1,2,4-triazol-3-yl, 4-oxo-2-benzopyranyl;

Q is COOH,

one of X^1 , X^2 , or X^3 or X^4 is nitrogen and the others are independently selected from CH and C-R_g and R_g is selected from 1) C₁₋₆alkyl optionally substituted with one to eight groups independently selected from aryl, heteroaryl, halogen, NR^aR^b, C(O)R^a, C(OR^a)R^aR^b, SR^a and OR^a, wherein aryl, heteroaryl and alkyl are each optionally substituted with one to six groups independently selected from halogen, CF₃, and COOH, or 2) S(O)_nC₁₋₆alkyl, wherein alkyl is optionally substituted with one to six substituents selected from halogen, aryl, heteroaryl, OH, and OC(O)R^a;

X² is CH;

X⁴ is CH or C-R_g, where R_g is selected from 1) C₁₋₆alkyl optionally substituted with OR^a or 2) S(O)_nC₁₋₆alkyl;

Y¹ is S;

Y² is selected from (CR^dRe)_m and CR^d=CR^e;

R¹ is selected from H, CN, OR^a, S(O)_nC₁₋₆alkyl and C₁₋₆alkyl optionally substituted with one to six groups independently selected from halogen, OR^a and S(O)_nC₁₋₆alkyl;

R² is selected from H and C₁₋₆alkyl optionally substituted with one to six halogen; or

R³ is selected from H and C₁₋₆alkyl optionally substituted with one to six groups independently selected from OR^a and halogen;

R^a and R^b are is independently selected from H; and C₁₋₁₀alkyl, C₂₋₁₀alkenyl, C₂₋₁₀alkynyl, Cy and Cy-C₁₋₁₀alkyl, wherein said alkyl, alkenyl, alkynyl and Cy are optionally substituted with one to six substituents independently selected from halogen, amino, carboxy, C₁₋₄alkyl, C₁₋₄alkoxy, aryl, heteroaryl, aryl-C₁₋₄alkyl, hydroxy, CF₃, OC(O)C₁₋₄alkyl, OC(O)NRⁱR^j, and aryloxy; or

R^e is selected from C₁₋₆alkyl optionally substituted with one to six halogen, aryl and heteroaryl, wherein said aryl and heteroaryl are optionally substituted with one to three groups selected from halogen, OC₁₋₆alkyl, O-haloC₁₋₆alkyl, C₁₋₆alkyl and haloC₁₋₆alkyl;

R^d and R^e are independently H, halogen, aryl, heteroaryl, C₁₋₆alkyl or haloC₁₋₆alkyl;

R^f is selected from H, C₁₋₆alkyl, haloC₁₋₆alkyl, Cy, C(O)C₁₋₆alkyl, C(O)haloC₁₋₆alkyl, and C(O)-Cy;

R_g is selected from

(1) — halogen,

(2) — CN,

(3) — C₁₋₆alkyl optionally substituted with one to eight groups

independently selected from aryl, heteroaryl, halogen, NR^aR^b,

C(O)R^a, C(OR^a)R^aR^b, SR^a and OR^a, wherein aryl, heteroaryl and

- ~~alkyl are each optionally substituted with one to six groups independently selected from halogen, CF₃, and COOH;~~
- ~~(4) — C₂₋₆alkenyl optionally substituted with one to six groups independently selected from halogen and OR^a;~~
- ~~(5) — Cy~~
- ~~(6) — C(O)R^a;~~
- ~~(7) — C(O)OR^a;~~
- ~~(8) — CONR^aR^b;~~
- ~~(9) — OCONR^aR^b;~~
- ~~(10) — OC₁₋₆alkyl, wherein alkyl is optionally substituted with one to six substituents selected from halogen, aryl, heteroaryl, OH and OC(O)R^a;~~
- ~~(11) — O—Cy;~~
- ~~(12) — S(O)_mC₁₋₆alkyl, wherein alkyl is optionally substituted with one to six substituents selected from halogen, aryl, heteroaryl, OH, and OC(O)R^a;~~
- ~~(13) — S(O)_m—Cy;~~
- ~~(14) — NR^aS(O)_mR^b;~~
- ~~(15) — NR^aR^b;~~
- ~~(16) — NR^aC(O)R^b;~~
- ~~(17) — NR^aC(O)OR^b;~~
- ~~(18) — NR^aC(O)NR^aR^b;~~
- ~~(19) — S(O)_mNR^aR^b;~~
- ~~(20) — NO₂;~~
- ~~(21) — C₅₋₈cycloalkenyl;~~

~~wherein Cy is optionally substituted with one to eight groups independently selected from halogen, C(O)R^a, OR^a, C₁₋₃alkyl, aryl, heteroaryl and CF₃;~~

~~Rⁱ and R^j are independently selected from hydrogen, C₁₋₁₀alkyl, Cy and Cy-C₁₋₁₀alkyl; or Rⁱ and R^j together with the nitrogen atom to which they are attached form a ring of 5 to 7 members containing 0-2 additional heteroatoms independently selected from oxygen, sulfur and N-R^f;~~

~~Cy is selected from heterocyclyl, aryl, and heteroaryl;~~

~~m is 1 or 2; and~~

~~n is 0, 1 or 2.~~

2. (Original) A compound of Claim 1 wherein A-Q is CH₂CO₂H.

3. (Cancel)

4. (Previously Canceled)

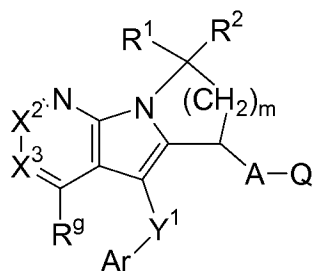
5. (Previously Canceled)

6. (Amended) A compound of Claim 1 wherein one of X^1 , ~~X^2~~ and X^3 is nitrogen and the other is ~~others are~~ CH, X^2 is CH, and X^4 is C-S(O)_n-C₁₋₆alkyl or C-C₁₋₆alkyl optionally substituted with OR^a.

7. (Cancel)

8. (Original) A compound of Claim 1 wherein Y^2 is selected from CH₂ and CH₂CH₂.

9. (Original) A compound of Claim 1 represented by the formula Ia:



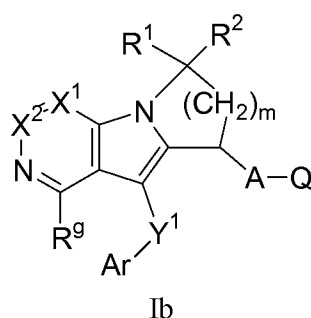
Ia

wherein X^2 and X^3 are independently CH or C-R_g, A, Ar, Q, Y^1 , R¹, R², m and R_g are as defined in Claim 1.

10. (Original) A compound of Claim 9 wherein X^2 and X^3 are each CH, R¹ and R² are each H, and A-Q is CH₂CO₂H.

11. (Original) A compound of Claim 9 wherein Y^1 -Ar is S-phenyl optionally substituted with 1 or 2 groups independently selected from halogen, C₁₋₆ alkyl and trifluoromethyl.

12. (Original) A compound of Claim 1 represented by the formula Ib:

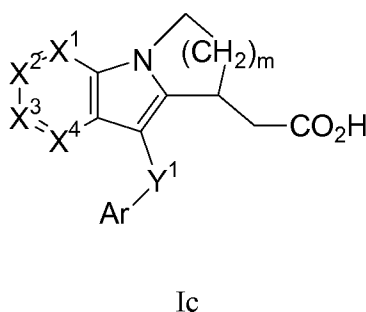


wherein X¹ and X² are independently CH or C-R_g, A, Ar, Q, Y¹, R¹, R², m and R_g are as defined in Claim 1.

13. (Original) A compound of Claim 12 wherein X¹ and X² are each CH, R¹ and R² are each H, and A-Q is CH₂CO₂H.

14. (Original) A compound of Claim 13 wherein Y¹-Ar is S-phenyl optionally substituted with 1 or 2 groups independently selected from halogen, C₁₋₆ alkyl and trifluoromethyl.

15. (Amended) A compound of Claim 1 represented by the formula Ic:



wherein one of X¹, ~~X²~~ and X³ is N and the others ~~are each~~ is CH, X⁴ is CH, X⁴ is CR_g, m is 1 or 2, and Ar, Y¹ and m are as defined in Claim 1.

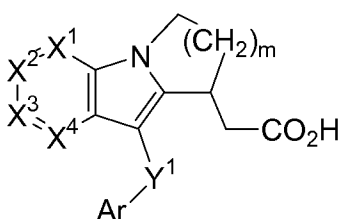
16. (Original) A compound of Claim 15 wherein Ar is phenyl optionally substituted with 1 or 2 groups independently selected from halogen, C₁₋₃alkyl and trifluoromethyl.

17. (Previously Canceled)

18. (Original) A compound of Claim 15 wherein X⁴ is selected from C-S(O)_n-C₁₋₆alkyl and C-C₁₋₆alkyl optionally substituted with OR^a.

19. (Previously Amended) A compound of Claim 15 wherein Y¹-Ar is S-phenyl optionally substituted with 1 or 2 groups independently selected from halogen, C₁₋₆alkyl and trifluoromethyl; X¹ and X² are each CH, X³ is N, m is 1 or 2, and X⁴ is C-SO₂C₁₋₆alkyl or C-C₁₋₆alkyl.

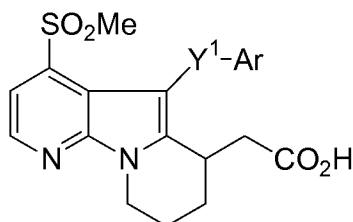
20. (Twice Amended) A compound of Claim 1 selected from:



X ¹	X ²	X ³	X ⁴	Ar	Y ¹	m
N	CH	CH	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
N	CH	CH	C(SCH ₃)	4-Cl-Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	3,4-diCl-Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	4-Br-Ph	S	2
CH	CH	N	C(SO ₂ CH ₃)	3,4-diCl-Ph	S	1
CH	CH	N	C(SO ₂ CH ₃)	3,4-diCl-Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	4-CF ₃ -Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	2-Cl-4-F-Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	2-naphthyl	S	2
N	CH	CH	C(SO ₂ CH ₃)	2,3-diCl-Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	4-CH ₃ -Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	Ph	S	2
N	CH	CH	C(SO ₂ CH ₃)	2,4-diCl-Ph	S	2
CH	N	CH	C(SO₂CH₃)	4-Cl-Ph	S	2
CH	CH	N	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
N	C(CH ₃)	CH	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
N	CH	C(CH ₃)	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
CH	C(CH ₃)	N	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
C(CH ₃)	CH	N	C(SO ₂ CH ₃)	4-Cl-Ph	S	2
N	CH	CH	C(CH(CH ₃) ₂)	4-F-Ph	S	2

X1	X2	X3	X4	Ar	Y1	m
N	CH	CH	C(CH(CH ₃) ₂)	4-Cl-Ph	S	2
N	CH	CH	C(CH(CH ₃) ₂)	2,4-diCl-Ph	S	2
N	CH	CH	C(CH(CH ₃) ₂)	4-Br-Ph	S	2
N	CH	CH	C(CH(CH ₃) ₂)	2-Cl-4-F-Ph	S	2
N	CH	CH	C(CH(CH ₃) ₂)	3,4-diCl-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	4-F-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	4-Cl-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	2,4-diCl-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	4-Br-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	2-Cl-4-F-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	3,4-diCl-Ph	S	2
CH	CH	N	C(CH(CH ₃) ₂)	4-F-Ph	S	1
CH	CH	N	C(CH(CH ₃) ₂)	4-Cl-Ph	S	1
CH	CH	N	C(CH(CH ₃) ₂)	2,4-diCl-Ph	S	1
CH	CH	N	C(CH(CH ₃) ₂)	4-Br-Ph	S	1
CH	CH	N	C(CH(CH ₃) ₂)	2-Cl-4-F-Ph	S	1
CH	CH	N	C(CH(CH ₃) ₂)	3,4-diCl-Ph	S	1
CH	N	CH	C(CH(CH ₃) ₂)	4-F-Ph	S	1
CH	N	CH	C(CH(CH ₃) ₂)	4-Cl-Ph	S	1
CH	N	CH	C(CH(CH ₃) ₂)	2,4-diCl-Ph	S	1
CH	N	CH	C(CH(CH ₃) ₂)	4-Br-Ph	S	1
CH	N	CH	C(CH(CH ₃) ₂)	2-Cl-4-F-Ph	S	1
CH	N	CH	C(CH(CH ₃) ₂)	3,4-diCl-Ph	S	1
CH	N	CH	C(CH(CH ₃) ₂)	4-F-Ph	S	2
CH	N	CH	C(CH(CH ₃) ₂)	4-Cl-Ph	S	2
CH	N	CH	C(CH(CH ₃) ₂)	2,4-diCl-Ph	S	2
CH	N	CH	C(CH(CH ₃) ₂)	4-Br-Ph	S	2
CH	N	CH	C(CH(CH ₃) ₂)	2-Cl-4-F-Ph	S	2
CH	N	CH	C(CH(CH ₃) ₂)	3,4-diCl-Ph	S	2
N	CH	CH	C(CH(OCH ₃)(CH ₂ CH ₃))	4-Cl-Ph	S	2
N	CH	CH	C(CH(OCH ₃)(CH ₂ CH ₃))	4-Cl-Ph	S	1
CH	N	CH	C(CH(OCH ₃)(CH ₂ CH ₃))	4-Cl-Ph	S	1

X1	X2	X3	X4	Ar	Y1	m
CH	N	CH	C(CH(OCH ₃))(CH ₂ CH ₃)	4-Cl-Ph	S	2
CH	CH	N	C(CH(OCH ₃))(CH ₂ CH ₃)	4-Cl-Ph	S	2
CH	CH	N	C(CH(OCH ₃))(CH ₂ CH ₃)	4-Cl-Ph	S	1
N	CH	CH	C(C(CH ₃) ₃)	4-Cl-Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	3,4-diCl-Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	4-Br-Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	4-CF ₃ -Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	2-Cl-4-F-Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	2-naphthyl	S	2
N	CH	CH	C(C(CH ₃) ₃)	2,3-diCl-Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	4-CH ₃ -Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	Ph	S	2
N	CH	CH	C(C(CH ₃) ₃)	2,4-diCl-Ph	S	2



Ar	Y1
5-tetrazolyl	S
2-pyrrolyl	S
1,2,4-triazol-3-yl	S
1,2,3-triazol-4-yl	S
5-imidazolyl	S
4-pyrazolyl	S
5-pyrazolyl	S
(1H,4H)-5-oxo-1,2,4-triazol-3-yl	S
4-isothiazolyl	S
1,2,5-thiadiazol-5-yl	S
1,2,5-oxadiazol-5-yl	S

Ar	Y1
3-furanyl	S
1,2,3-thiadiazol-4-yl	S
1,2,3-oxadiazol-4-yl	S
4-isoxazolyl	S
3-thienyl	S
4-oxazolyl	S
4-thiazolyl	S
(5H)-2-oxo-5-furanyl	S
(5H)-2-oxo-4-furanyl	S
1,2,4-oxadiazol-5-yl	S
3-pyridyl	S
2-pyrazinyl	S
5-pyrimidinyl	S
2-indolyl	S
2-benzothienyl	S
2-benzofuranyl	S
4-oxo-benzopyran-2-yl	S
2-quinoliny	S
2-benzimidazolyl	S
2-benzoxazolyl	S
2-benzothiazolyl	S
1-benzotriazolyl	CH ₂ S
thieno[2,3-b]pyridin-2-yl	S

21. (Original) A pharmaceutical composition comprising a compound of Claim 1 and a pharmaceutically acceptable carrier.

22. (Cancel)

23. (Cancel)

24. (Cancel)

25. (Cancel)

26. (Cancel)

27. (Previously Canceled)
28. (Previously Canceled)
29. (Previously Canceled)
30. (Previously Canceled)